

AMENDED VERSION

IN THE CLAIMS:

SUB  
E1

D1

5. <sup>Twice</sup>  
(Amended)

A synthetic nuclease resistant antisense oligodeoxynucleotide for selectively inhibiting human tumor necrosis factor alpha said antisense oligonucleotide comprising: an exon targeting a sequence which flanks at least one splice site said targeting thereby regulating expression of TNF- $\alpha$ .

D2

7. <sup>Thrice</sup>  
(Twice Amended)

A pharmaceutical composition for selectively inhibiting mammalian tumor necrosis factor alpha in a mammal in need of such treatment consisting of

an effective amount of at least one active ingredient a synthetic nuclease resistant antisense oligodeoxynucleotide having a nucleotide sequence selected from the group consisting of SEQ. ID No. 4 and SEQ. ID No. 6 in a pharmaceutically physiologically acceptable carrier or diluent.

D3

13. <sup>Twice</sup>  
(Amended)

A method of selectively regulating mammalian tumor necrosis factor alpha by targeting for treatment a tumor necrosis factor alpha splice region and then specifically modify the region to inhibit the mammalian tumor necrosis factor alpha.

D4

14. <sup>Twice</sup>  
(Amended)

The method of claim 13 further including administering an effective amount of a synthetic nuclease resistant antisense oligodeoxynucleotide which targets exon sequences flanking donor splice sites.

15.<sup>Twice</sup> (Amended) A method of inhibiting tumor necrosis factor alpha by targeting for treatment a tumor necrosis factor alpha splice region thereby inhibiting tumor necrosis factor alpha.

DS

16.<sup>Twice</sup> (Amended) The method of claim 15 further including administering an effective amount of a synthetic nuclease resistant antisense oligodeoxynucleotide which targets exon sequences flanking donor splice sites.